AGGCGGCAGCAGCTGCAGGCTGACCTTGCAGCTTGGCGGAATGGACTGGCCTCACAACCTGCTGTTTCTT
CTTACCATTTCCATCTTCCTGGGGCTGGGCCAGCCCAGGAGCCCCAAAAGCAAGAGGAAGGGGCAAGGGCG
GCCTGGGCCCTGGCCCTCACCAGGTGCCACTGGACCTGGTGTCACGGATGAAACCGTATGCCC
GCATGGAGGAGTATGAGAGAGAACATCGAGGAGATGGTGGCCCAGCTGAGGAACAGCTCAGAGCTGGCCCAG
AGAAAGTGTGAGGTCAACTTGCAGCTGTGGATGTCCAACAAGAGGAGCCTGTCTCCCTGGGGCTACAGCAT
CAACCACGACCCCAGCCGTATCCCCGTGGACCTGCCGGAGGCACGGTGCCTGTGTCTCGGCCGCCGC
CCTTCACCATGCAGGAGACCGCAGCATGGTGAGCGTGCCGGTGTTCAGCCAGGTTCCTGTGCGCCGCCGC
CTCTGCCCGCCACCGCCCCCCCCACAGGGCCTTGCCGCCAGCGCCAGCAGCACCATCCTTTGCACCTTT
CACCTGCATCTTCTGAATCACCTGGCCCAGAAGCCAGGCCAGCAGCCCGAGACCATCCTCTTTGCACCTTT
GTGCCCAAGAAAGGCCTATGAAAAGTAAACACTGACTTTTGAAAGCAAG

FIGURE 2

 ${\tt MDWPHNLLFLLTISIFLGLGQPRSPKSKRKGQGRPGPLAPGPHQVPLDLVSRMKPYARMEEYERNIEEMVA} \\ QLRNSSELAQRKCEVNLQLWMSNKRSLSPWGYSINHDPSRIPVDLPEARCLCLGCVNPFTMQEDRSMVSVP\\ VFSQVPVRRRLCPPPPRTGPCRQRAVMETIAVGCTCIF$

FIGURE 4

Signal peptide:

Tyrosine kinase phosphorylation site:

N-myristoylation sites:

Leucine zipper pattern:

Homologous region to IL-17:

Amino acids 1-18

Amino acids 112-121

Amino acids

32-38;55-61;133-139

Amino acids 3-25

Amino acids 99-195

GTGCCCGACTTGTGACTGAGTGTGCAGTGCCCAGCATGTACCAGGTCAGTGCAGAGGGCTGCCTGAGGGCT GTGCTGAGAGGGGGGGGGGGAGGCAGGATGCTGCTGAGGGTTGGAGGGGAGGCCAAGCTGCCAGGTTTGGGGCTGG $\tt GGGCCAAGTGGAGTGAGAAACTGGGATCCCAGGGGGAGGGTGCAG\underline{ATG} AGGGAGCGACCCAGATTAGGTGA$ ${\tt GGACAGTTCTCATTAGCCTTTTCCTACAGGTGGTTGCATTCTTGGCAATGGTCATGGGAACCCACACCT}$ ACAGCCACTGGCCCAGCTGCCCCCAGCAAAGGGCAGGACACCTCTGAGGAGCTGCTGAGGTGGAGCACT GTGCCTGTGCCTCCCCTAGAGCCTGCTAGGCCCAACCGCCACCCAGAGTCCTGTAGGGCCAGTGAAGATGGA $\tt CCCCTCAACAGCAGGCCCATCTCCCCCTGGAGATATGAGTTGGACAGAGACTTGAACCGGCTCCCCCAGGA$ $\tt GCAACTCGGAGCTGCTCTACCACAACCAGACTGTCTTCTACAGGCGGCCATGCCATGGCGAGAAGGGCACC$ GATGGGCTAGCCGGACCTGCTGGAGGCTGGTCCCTTTTTGGGAAACCTGGAGCCAGGTGTACAACCACTTG AGGATGGGGGGCTTTGGGGAAAACCTGCACTTCTGCACATTTTGAAAAGAGCAGCTGCTGCTTAGGGCCGC $\tt CGGAAGCTGGTGTCCTGTCATTTTCTCTCAGGAAAGGTTTTCAAAGTTCTGCCCATTTCTGGAGGCCACCA$ $\tt CTCCTGTCTCTTTCCCATCCCCTGCTACCCTGGCCCAGCACAGGCACTTTCTAGATATTTCCCC$ GTGCATTCTAGTGTAGTTACTAGTCTTTTGACATGGATGATTCTGAGGAGGAAGCTGTTATTGAATGTATA GAGATTTATCCAAATAAATATCTTTATTTAAAAATGAAAA

 ${\tt MRERPRLGEDSSLISLFLQVVAFLAMVMGTHTYSHWPSCCPSKGQDTSEELLRWSTVPVPPLEPARPNRHP}$ ESCRASEDGPLNSRAISPWRYELDRDLNRLPQDLYHARCLCPHCVSLQTGSHMDPRGNSELLYHNQTVFYR RPCHGEKGTHKGYCLERRLYRVSLACVCVRPRVMG

Signal peptide:

Amino acids 1-32

N-glycosylation site:

Amino acids 136-140

Tyrosine kinase phosphorylation site: Amino acids 127-135

N-myristoylation sites:

Amino acids 44-50;150-156

ATGCTGGTAGCCGGCTTCCTGCTGCGCCCGAGCTGGGCCGCGGGCGCCCCCAGGCCGGCAGGC CCCGCGCGCCGCGGGCTGCGCGGACCGGCCGGAGGAGCTACTGGAGCAGCTGTACGGGCGCCTGGCGG CCGGCGTGCTCAGTGCCTTCCACCACACGCTGCAGCTGGGGCCGCGTGAGCAGGCGCGCAACGCGAGCTGC CCGGCAGGGGCAGGCCCGGCGACCGCCGCTTCCGGCCGCCCACCAACCTGCGCAGCGTGTCGCCCTGGGC ${\tt GCCTGACCGGGCTGTTCGGCGAGGAGGAGGACGTGCGCTTCCGCAGCGCCCCTGTCTACATGCCCACCGTCGTC}$ $\tt CTGCGCCGCACCCCGCCTGCGCCGGCGGCCGTTCCGTTACACCGAGGCCTACGTCACCATCCCCGTGGG$ CTGCACCTGCGTCCCCGAGCCGGAGAAGGACGCAGACATCAACTCCAGCATCGACAAACAGGGCGCCA AGCTCCTGCTGGGCCCCAACGACGCCCCGCTGGCCCCTGAGGCCGGTCCTGCCCCGGGAGGTCTCCCCGG ${\tt GAGCAAACCAAGTGCCGGAGCACCAGCGCCGCCTTTCCATGGAGACTCGTAAGCAGCTTCATCTGACACGG}$ GCATCCCTGGCTTGCTTTTAGCTACAAGCAAGCAGCGTGGCTGGAAGCTGATGGGAAACGACCCGGCACGG GCATCCTGTGTGCGGCCCGCATGGAGGGTTTGGAAAAGTTCACGGAGGCTCCCTGAGGAGCCTCTCAGATC $\tt GGCTGCTGCGGGTGCAGGGCGTGACTCACCGCTGGGTGCTTGCCAAAGAGATAGGGACGCATATGCTTTTT$ AAAGCAATCTAAAAATAATAATAAGTATAGCGACTATATACCTACTTTTAAAATCAACTGTTTTGAATAGA ${\tt GGCAGAGCTATTTTATATTATCAAATGAGAGCTACTCTGTTACATTTCTTAACATATAAACATCGTTTTTT}$ ACTTCTTCTGGTAGAATTTTTTAAAGCATAATTGGAATCCTTGGATAAATTTTGTAGCTGGTACACTCTGG TTCCTTCCACTGAAGGTCTTCACGGGCCTCCAGGTGGACCAAAGGGATGCACAGGCGGCTCGCATGCCCCA $\tt GGGCCAGCTAAGAGTTCCAAAGATCTCAGATTTGGTTTTAGTCATGAATACATAAACAGTCTCAAACTCGC$ ACAATTTTTTCCCCCTTTTGAAAGCCACTGGGGCCAATTTGTGGTTAAGAGGTGGTGAGATAAGAAGTGGA ${\tt ACGTGACATCTTTGCCAGTTGTCAGAAGAATCCAAGCAGGTATTGGCTTAGTTGTAAGGGCTTTAGGATCA}$ GGCTGAATATGAGGACAAAGTGGGCCACGTTAGCATCTGCAGAGATCAATCTGGAGGCTTCTGTTTCTGCA AAAAGTTAGTAGTTCTTTTTAAATCATTAAAAGAGGCTTGCTGAAGGAT

Signal peptide:

Amino acids 1-15

N-glycosylation sites:

Amino acids 68-72;181-185

Tyrosine kinase phosphorylation site:

Amino acids 97-106

N-myristoylation sites:

Amino acids 17-23;49-55;74-80;

118-124

Amidation site:

Amino acids 21-25

FIGURE 9

FIGURE 10

 ${\tt MTVKTLHGPAMVKYLLLSILGLAFLSEAAARKIPKVGHTFFQKPESCPPVPGGSMKLDIGIINENQRVSMS} \\ RNIESRSTSPWNYTVTWDPNRYPSEVVQAQCRNLGCINAQGKEDISMNSVPIQQETLVVRRKHQGCSVSFQ\\ LEKVLVTVGCTCVTPVIHHVQ\\$

Signal sequence:

Amino acids 1-30

N-glycosylation site:

Amino acids 83-86

N-myristoylation sites:

Amino acids 106-111;136-141

FIGURE 11

 $\tt CCGGCGATGTCGCTGCTGCTAAGCCTGGCCGCTGTGCAGGAGCGCCGTACCCCGAGAGCCGACCGT$ TCAATGTGGCTCTGAAACTGGGCCATCTCCAGAGTGGATGCTACAACATGATCTAATCCCCGGAGACTTGA GGGACCTCCGAGTAGAACCTGTTACAACTAGTGTTGCAACAGGGGACTATTCAATTTTGATGAATGTAAGC TGGGTACTCCGGGCAGATGCCAGCATCCGCTTGTTGAAGGCCACCAAGATTTGTGTGACGGGCAAAAGCAA CTTCCAGTCCTACAGCTGTGTGAGGTGCAATTACACAGAGGCCTTCCAGACTCAGACCCAGACCCTCTGGTG $\tt GTAAATGGACATTTTCCTACATCGGCTTCCCTGTAGAGCTGAACACAGTCTATTTCATTGGGGCCCATAAT$ ATTCCTAATGCAAATATGAATGAAGATGGCCCTTCCATGTCTGTGAATTTCACCTCACCAGGCTGCCTAGA CCACATAATGAAATATAAAAAAAAGTGTGTCAAGGCCGGAAGCCTGTGGGGATCCGAACATCACTGCTTGTA AGAAGAATGAGGAGACAGTAGAAGTGAACTTCACAACCACTCCCCTGGGAAACAGATACATGGCTCTTATC GGTGATTCCAGTGACTGGGGATAGTGAAGGTGCTACGGTGCAGCTGACTCCATATTTTCCTACTTGTGGCA GCGACTGCATCCGACATAAAGGAACAGTTGTGCTCTGCCCACAAACAGGCGTCCCTTTCCCTCTGGATAAC AACAAAAGCAAGCCGGGAGGCTGGCTGCCTCTCCTCCTGCTGCTGCTGGTGGCCACATGGGTGCTGGT GGCAGGGATCTATCTAATGTGGAGGCACGAAAGGATCAAGAAGACTTCCTTTTCTACCACCACACTACTGC $\tt CCCCCATTAAGGTTCTTGTGGTTTACCCATCTGAAATATGTTTCCATCACACAATTTGTTACTTCACTGAA$ TTTCTTCAAAACCATTGCAGAAGTGAGGTCATCCTTGAAAAGTGGCAGAAAAAGAAAATAGCAGAGATGGG TCCAGTGCAGTGGCTTGCCACTCAAAAGAAGGCAGCAGACAAAGTCGTCTTCCTTTCCAATGACGTCA ${\tt ACAGTGTGCGATGGTACCTGTGGCAAGAGCGAGGGCAGTCCCAGTGAGAACTCTCAAGACCTCTTCCCCC}$ CTTGCCTTTAACCTTTTCTGCAGTGATCTAAGAAGCCAGATTCATCTGCACAAATACGTGGTGGTCTACTT TAGAGAGATTGATACAAAAGACGATTACAATGCTCTCAGTGTCTGCCCCAAGTACCACCTCATGAAGGATG CCACTGCTTTCTGTGCAGAACTTCTCCATGTCAAGCAGCAGGTGTCAGCAGGAAAAAAGATCACAAGCCTGC CACGATGGCTGCTGCTCCTTGTAG

FIGURE 12

MSLVLLSLAALCRSAVPREPTVQCGSETGPSPEWMLQHDLIPGDLRDLRVEPVTTSVATGDYSILMNVSWV LRADASIRLLKATKICVTGKSNFQSYSCVRCNYTEAFQTQTRPSGGKWTFSYIGFPVELNTVYFIGAHNIP NANMNEDGPSMSVNFTSPGCLDHIMKYKKKCVKAGSLWDPNITACKKNEETVEVNFTTTPLGNRYMALIQH STIIGFSQVFEPHQKKQTRASVVIPVTGDSEGATVQLTPYFPTCGSDCIRHKGTVVLCPQTGVPFPLDNNK SKPGGWLPLLLLSLLVATWVLVAGIYLMWRHERIKKTSFSTTTLLPPIKVLVVYPSEICFHHTICYFTEFL QNHCRSEVILEKWQKKKIAEMGPVQWLATQKKAADKVVFLLSNDVNSVCDGTCGKSEGSPSENSQDLFPLA FNLFCSDLRSQIHLHKYVVVYFREIDTKDDYNALSVCPKYHLMKDATAFCAELLHVKQQVSAGKRSQACHD GCCSL

Signal sequence:

Amino acids 1-14

Transmembrane domain:

Amino acids 290-309

N-glycosylation sites:

Amino acids 67-70;103-106;156-159;

183-186;197-200;283-286

cAMP- and cGMP-dependent protein kinase phosphorylation sites:

Amino acids 228-231;319-322

N-myristoylation site:

Amino acids 116-121

Amidation site:

Amino acids 488-491

FIGURE 13

ACACTGGCCAAACAAAAACGAAAGCACTCCGTGCTGGAAGTAGGAGGAGAGTCAGGACTCCCAGGACAGAG AGTGCACAAACTACCCAGCACAGCCCCCTCCGCCCCCTCTGGAGGGCTGAAGAGGGGATTCCAGCCCCTGCCA GCCACCTGGCACCTAGAAG ATG CCTGTGCCCTGGTTCTTGCTGTCCTTGGCACTGGGCCGAAGCCCAGTGGTCCTTTCTCTGGAGAGGCTTGTGGGGCCTCAGGACGCTACCCACTGCTCTCCGGGCCTCTCCTGCCGCCTC TGGGACAGTGACATACTCTGCCTGCCTGGGGACATCGTGCCTGCTCCGGGCCCCGTGCTGGCGCCTACGCA ACTTGGCCGTGCATGGGCACTGGGAAGACCCTGAAGATGAGGAAAAGTTTGGAGGAGCAGCTGACTCAGGG GTGGAGGAGCCTAGGAATGCCTCTCCCAGGCCCAAGTCGTGCTCTCCTTCCAGGCCTACCCTACTGCCCG $\tt CTGCGTCCTGCTGGAGGTGCAAGTGCCTGCTGCCCTTGTGCAGTTTGGTCAGTCTGTGGGCTCTGTGGTATAT$ GACTGCTTCGAGGCTGCCCTAGGGAGTGAGGTACGAATCTGGTCCTATACTCAGCCCAGGTACGAGAAGGA ACTCAACCACACACAGCAGCTGCCTGCCCTGCCCTGGCTCAACGTGTCAGCAGATGGTGACAACGTGCATC TGGTTCTGAATGTCTCTGAGGAGCAGCACTTCGGCCTCTCCCTGTACTGGAATCAGGTCCAGGGCCCCCCA CCTCTGTATTCAGGTGTGGCCTCTGGAACCTGACTCCGTTAGGACGAACATCTGCCCCTTCAGGGAGGACC GGTCCCACCGCTTTCCTGGGAGAACGTCACTGTGGACAAGGTTCTCGAGTTCCCATTGCTGAAAGGCCACC CTAACCTCTGTGTTCAGGTGAACAGCTCGGAGAAGCTGCAGCTGCAGGAGTGCTTGTGGGCTGACTCCCTG GGGCCTCTCAAAGACGATGTGCTACTGTTGGAGACACGAGGCCCCCAGGACAACAGATCCCTCTGTGCCTT GGAACCCAGTGGCTGTACTTCACTACCCAGCAAAGCCTCCACGAGGGCAGCTCGCCTTGGAGAGTACTTAC TACAAGACCTGCAGTCAGGCCAGTGTCTGCAGCTATGGGACGATGACTTGGGAGCGCTATGGGCCTGCCCC CCTCATCCTCCTTCTCAAAAAGGATCACGCGAAAGGGTGGCTGAGGCTCTTGAAACAGGACGTCCGCTCGG GGGCGCCCCAGGGGCCGCGCGCTCTGCTCCTCTACTCAGCCGATGACTCGGGTTTCGAGCGCCTGGTG GGCGCCTGGCGTCGGCCCTGTGCCAGCTGCCGCTGCGCGTGGCCGTAGACCTGTGGAGCCGTCGTGAACT GAGCGCGCAGGGGCCCGTGGCTTGGTTTCACGCGCAGCGGCGCCAGACCCTGCAGGAGGGCGGCGTGGTGG TCTTGCTCTTCTCCCGGTGCGGTGGCGCTGTGCAGCGAGTGGCTACAGGATGGGGTGTCCGGGCCCGGG GCGCACGGCCCGCACGACGCCTTCCGCGCCTCGCTCAGCTGCTGCCCCGACTTCTTGCAGGGCCGGGC GCCCGCCAGCTACGTGGGGGCCTGCTTCGACAGGCTGCTCCACCCGGACGCCGTACCCGCCCTTTTCCGCA CCGTGCCGTCTTCACACTGCCCCAACTGCCAGACTTCCTGGGGGCCCTGCAGCAGCCTCGCGCCCCG CCATCCCCGGGGACTCCCGCGCGGGACGCGGGGGGGACCAGGGGCGGGACCTGGGGCGGGGACGGGA CTTAAATAAAGGCAGACGCTGTTTTTCTAAAAAAA

FIGURE 14

MPVPWFLLSLALGRSPVVLSLERLVGPQDATHCSPGLSCRLWDSDILCLPGDIVPAPGPVLAPTHLQTELV $\tt LRCQKETDCDLCLRVAVHLAVHGHWEEPEDEEKFGGAADSGVEEPRNASLQAQVVLSFQAYPTARCVLLEV$ ${\tt QVPAALVQFGQSVGSVVYDCFEAALGSEVRIWSYTQPRYEKELNHTQQLPALPWLNVSADGDNVHLVLNVS}$ EEQHFGLSLYWNQVQGPPKPRWHKNLTGPQIITLNHTDLVPCLCIQVWPLEPDSVRTNICPFREDPRAHQN LWOAARLRLLTLOSWLLDAPCSLPAEAALCWRAPGGDPCQPLVPPLSWENVTVDKVLEFPLLKGHPNLCVQ VNSSEKLOLOECLWADSLGPLKDDVLLLETRGPQDNRSLCALEPSGCTSLPSKASTRAARLGEYLLQDLQS GQCLQLWDDDLGALWACPMDKYIHKRWALVWLACLLFAAALSLILLLKKDHAKGWLRLLKQDVRSGAAARG ${\tt RAALLLYSADDSGFERLVGALASALCQLPLRVAVDLWSRRELSAQGPVAWFHAQRRQTLQEGGVVVLLFSP}$ GAVALCSEWLQDGVSGPGAHGPHDAFRASLSCVLPDFLQGRAPGSYVGACFDRLLHPDAVPALFRTVPVFT $\verb|LPSQLPDFLGALQQPRAPRSGRLQERAEQVSRALQPALDSYFHPPGTPAPGRGVGPGAGPGAGDGT|$

signal sequence:

Amino acids 1-20

transmembrane domain:

Amino acids 453-473

N-glycosylation sites:

Amino acids 118-121;186-189;198-201;

211-214;238-241;248-251;334-337;

357-360;391-394

Glycosaminoglycan attachment site: Amino acids 583-586

cAMP- and cGMP-dependent protein kinase phosphorylation site:

Amino acids 552-555

N-myristoylation sites:

Amino acids 107-112;152-157;319-324;

438-443;516-521;612-617;692-697;

696-701;700-705

FIGURE 15

 $\tt CCATGTTCCGGGAGCCCTAATTGCACAGAAGCCCATGGGGAGCTCCAGACTGGCAGCCCTGCTCCTGCCTC$ TCCTCCTCATAGTCATCGACCTCTCTGACTCTGCTGGGATTGGCTTTCGCCACCTGCCCCACTGGAACACC CGCTGTCCTCTGGCCTCCCACACGGATGACAGTTTCACTGGAAGTTCTGCCTATATCCCTTGCCGCACCTG GTGGGCCCTCTTCTCCACAAAGCCTTGGTGTGTGCGAGTCTGGCACTGTTCCCGCTGTTTGTGCCAGCATCTGC TGTCAGGTGGCTCAGGTCTTCAACGGGGCCTCTTCCACCTCCTGGTGCAGAAAATCCAAAAAGTCTTCCACA $\tt TTCAAGTTCTATAGGAGACACAAGATGCCAGCACCTGCTCAGAGGAAGCTGCTGCTCGTCGTCACCTGTC$ TGAGAAGAGCCATCACATTTCCATCCCCTCCCCAGACATCTCCCACAAGGGACTTCGCTCTAAAAGGACCC AACCTTCGGATCCAGAGACATGGGAAAGTCTTCCCAGATTGGACTCACAAAGGCATGGAGGACCCGAGTTC TCCTTTGATTTGCTGCCTGAGGCCCGGGCTATTCGGGTGACCATATCTTCAGGCCCTGAGGTCAGCGTGCG TCTTTGTCACCAGTGGGCACTGGAGTGTGAAGAGCTGAGCAGTCCCTATGATGTCCAGAAAATTGTGTCTG GAGGACACTGTGAGGCGCAAAAAATGTCCCTTCCAGAGCTGGCCAGAAGCCTATGGCTCGGACTTCTGGAA GTCAGTGCACTTCACTGACTACAGCCAGCACACTCAGATGGTCATGGCCCTGACACTCCGCTGCCCACTGA AGCTGGAAGCTGCCCTCTGCCAGAGGCACGACTGGCATACCCTTTGCAAAGACCTCCCGAATGCCACGGCT $\tt CGAGAGTCAGATGGGTAGTTTTTGGAGAAGGTGGACCTGCACCCCAGCTCTGCTTCAAGTTCTCTTT$ TGGAAACAGCAGCCATGTTGAATGCCCCCACCAGACTGGGTCTCTCACATCCTGGAATGTAAGCATGGATA CCAGGCTTGGGGCAGGACACTTTGGTGCCCCCGTGTACACTGTCAGCCAGGCCCGGGGCTCAAGCCCAGT GTCACTAGACCTCATCATTCCCTTCCTGAGGCCAGGGTGCTGTGTCCTGGTGTGGCGGTCAGATGTCCAGT TTGCCTGGAAGCACCTCTTGTGTCCAGATGTCTCTTACAGACACCTGGGGCTCTTGATCCTGGCACTGCTG GCCCTCCTCACCCTACTGGGTGTTGTTCTGGCCCTCACCTGCCGGCCCCACAGTCAGGCCCGGGCCCAGC GCGGCCAGTGCTCCTCCTGCACGCGGCGGACTCGGAGGCGCAGCGGCGCCTGGTGGGAGCGCTGGAAC GGGCCCGCTGCCGTGGCTCTGGGCGCGCGCGCGCGCGTAGCGCGGGAGCAGGGCACTGTGCTGCTGCTGT $\tt GGAGCGGCCGACCTTCGCCCGGTCAGCGGCCCCGACCCCCGCGCGCCCCTGCTCGCCCTGCTCCAC$ ${\tt GCTGCCCGCGCCGCTGCTGCTGCTCGCTTACTTCAGTCGCCTCTGCGCCAAGGGCGACATCCCCCCGCC}$ GCTGCGCGCCCTGCCGCGTACCGCCTGCTGCGCGACCTGCCGCGTCTGCTGCGGGCGCTGGACCGCGCGC AGCCGGCTTGAACGAGAGGCCGCCCGACTTGCAGACCTAGGTTGAGCAGAGCTCCACCGCAGTCCCGGGTGTCT

FIGURE 16

MGSSRLAALLLPLLLIVIDLSDSAGIGFRHLPHWNTRCPLASHTDDSFTGSSAYIPCRTWWALFSTKPWCV RVWHCSRCLCQHLLSGGSGLQRGLFHLLVQKSKKSSTFKFYRRHKMPAPAQRKLLPRRHLSEKSHHISIPS PDISHKGLRSKRTQPSDPETWESLPRLDSQRHGGPEFSFDLLPEARAIRVTISSGPEVSVRLCHQWALECE ELSSPYDVQKIVSGGHTVELPYEFLLPCLCIEASYLQEDTVRRKKCPFQSWPEAYGSDFWKSVHFTDYSQH TOMVMALTLRCPLKLEAALCQRHDWHTLCKDLPNATARESDGWYVLEKVDLHPQLCFKFSFGNSSHVECPH OTGSLTSWNVSMDTOAOOLILHFSSRMHATFSAAWSLPGLGQDTLVPPVYTVSQARGSSPVSLDLIIPFLR ${\tt PGCCVLVWRSDVQFAWKHLLCPDVSYRHLGLLILALLALLTLLGVVLALTCRRPQSGPGPARPVLLLHAAD}$ SEAORRLVGALAELLRAALGGGRDVIVDLWEGRHVARVGPLPWLWAARTRVAREQGTVLLLWSGADLRPVS GPDPRAAPLLALLHAAPRPLLLLAYFSRLCAKGDIPPPLRALPRYRLLRDLPRLLRALDARPFAEATSWGR LGARQRRQSRLELCSRLEREAARLADLG

Signal peptide:

Amino acids 1-23

Transmembrane domain:

Amino acids 455-472

N-glycosylation sites:

Amino acids 318-322;347-351;364-368

Glycosaminoglycan attachment site: Amino acids 482-486

cAMP- and cGMP-dependent protein kinase phosphorylation sites:

Amino acids 104-108;645-649

Tyrosine kinase phosphorylation site:

Amino acids 322-329

N-myristoylation sites:

Amino acids 90-96;358-364;470-476

Eukaryotic cobalamin-binding proteins:

Amino acids 453-462

FIGURE 17

GAAACAGTGGGCTGTACAACATCACCTTCAAATATGACAATTGTACCACCTACTTGAATCCAGTGGGGAAG CATGTGATTGCTGACGCCCAGAATATCACCATCAGCCAGTATGCTTGCCATGACCAAGTGGCAGTCACCAT TCTTTGGTCCCCAGGGGCCCTCGGCATCGAATTCCTGAAAGGATTTCGGGTAATACTGGAGGAGCTGAAGT CGGAGGGAAGACAGTGCCAACAACTGATTCTAAAGGATCCGAAGCAGCTCAACAGTAGCTTCAAAAGAACT $\tt GGAATGGAATCTCAACCTTTCCTGAATATGAAATTTGAAACGGATTATTTCGTAAAGGTTGTCCCTTTTCC$ TTCCATTAAAAACGAAAGCAATTACCACCCTTTCTTCTTTAGAACCCGAGCCTGTGACCTGTTGTTACAGC AGGTGTCCTTCGACCACGCACGCATGGCTCGGACATGCAGGTGTCCTTCGACCACGCACCGCACAACTTC GGCTTCCGTTTCTTCTATCTTCACTACAAGCTCAAGCACGAAGGACCTTTCAAGCGAAAGACCTGTAAGCA GGAGCAAACTACAGAGATGACCAGCTGCCTCCTTCAAAATGTTTCTCCAGGGGATTATATAATTGAGCTGG TGGATGACACTAACACAACAAGAAAAGTGATGCATTATGCCTTAAAGCCAGTGCACTCCCCGTGGGCCGGG $\tt CCCATCAGAGCCGTGGCCATCACAGTGCCACTGGTAGTCATATCGGCATTCGCGACGCTCTTCACTGTGAT$ GTGCCGCAAGAAGCAACAAGAAAATATATATTCACATTTAGATGAAGAGAGCTCTGAGTCTTCCACATACA $\tt CTGCAGCACTCCCAAGAGAGGGCTCCGGCCGGCCGAAGGTCTTTCTCTGCTATTCCAGTAAAGATGGC$ CAGAATCACATGAATGTCGTCCAGTGTTTCGCCTACTTCCTCCAGGACTTCTGTGGCTGTGAGGTGGCTCT GGACCTGTGGGAAGACTTCAGCCTCTGTAGAGAAGGGCAGAGAAATGGGTCATCCAGAAGATCCACGAGT CCCAGTTCATCATTGTGGTTTGTTCCAAAGGTATGAAGTACTTTGTGGACAAGAAGAACTACAAACACAAA GGAGGTGGCCGAGGCTCGGGGAAAGGAGAGCTCTTCCTGGTGGCGGTGTCAGCCATTGCCGAAAAGCTCCG $\tt CCAGGCCAAGCAGAGTTCGTCCGCGGCGCTCAGCAAGTTTATCGCCGTCTACTTTGATTATTCCTGCGAGG$ GAGACGTCCCCGGTATCCTAGACCTGAGTACCAAGTACAGACTCATGGACAATCTTCCTCAGCTCTGTTCC CACCTGCACTCCCGAGACCACGGCCTCCAGGAGCCGGGGCAGCACACGCGACAGGGCAGCAGAAGGAACTA $\tt CTTCCGGAGCAAGTCAGGCCGGTCCCTATACGTCGCCATTTGCAACATGCACCAGTTTATTGACGAGGAGC$ CCGACTGGTTCGAAAAGCAGTTCGTTCCCTTCCATCCTCCACTGCGCTACCGGGAGCCAGTCTTGGAG AAATTTGATTCGGGCTTGGTTTTAAATGATGTCATGTGCAAACCAGGGCCTGAGAGTGACTTCTGCCTAAA GGTAGAGGCGGCTGTTCTTGGGGCAACCGGACCAGCCGACTCCCAGCACGAGAGTCAGCATGGGGGCCTGG ACCAAGACGGGGAGGCCCGGCCTGCCCTTGACGGTAGCGCCGCCCTGCAACCCCTGCTGCACACGGTGAAA $\tt GCCGGCAGCCCTCGGACATGCCGCGGGACTCAGGCATCTATGACTCGTCTGTGCCCTCATCCGAGCTGTC$ TCTGCCACTGATGGAAGGACTCTCGACGGACCAGACAGAAACGTCTTCCCTGACGGAGAGCGTGTCCTCCT GATCTTGGTTGCCGCAGCTACACTGATGAACTCCACGCGGTCGCCCCTTTGTAACAAAACGAAAGAGTCTA

FIGURE 18

MPRASASGVPALFVSGEQGVGPASRNSGLYNITFKYDNCTTYLNPVGKHVIADAQNITISQYACHDQVAVT
ILWSPGALGIEFLKGFRVILEELKSEGRQCQQLILKDPKQLNSSFKRTGMESQPFLNMKFETDYFVKVVPF
PSIKNESNYHPFFFRTRACDLLLQPDNLACKPFWKPRNLNISQHGSDMQVSFDHAPHGSDMQVSFDHAPHN
FGFRFFYLHYKLKHEGPFKRKTCKQEQTTEMTSCLLQNVSPGDYIIELVDDTNTTRKVMHYALKPVHSPWA
GPIRAVAITVPLVVISAFATLFTVMCRKKQQENIYSHLDEESSESSTYTAALPRERLRPRPKVFLCYSSKD
GQNHMNVVQCFAYFLQDFCGCEVALDLWEDFSLCREGQREWVIQKIHESQFIIVVCSKGMKYFVDKKNYKH
KGGGRGSGKGELFLVAVSAIAEKLRQAKQSSSAALSKFIAVYFDYSCEGDVPGILDLSTKYRLMDNLPQLC
SHLHSRDHGLQEPGQHTRQGSRRNYFRSKSGRSLYVAICNMHQFIDEEPDWFEKQFVPFHPPPLRYREPVL
EKFDSGLVLNDVMCKPGPESDFCLKVEAAVLGATGPADSQHESQHGGLDQDGEARPALDGSAALQPLLHTV
KAGSPSDMPRDSGIYDSSVPSSELSLPLMEGLSTDQTETSSLTESVSSSSGLGEEEPPALPSKLLSSGSCK
ADLGCRSYTDELHAVAPL

Transmembrane domain:

Amino acids 283-307

N-glycosylation sites:

Amino acids 31-34;38-41;56-59;

113-116;147-150;182-185;266-269

Glycosaminoglycan attachment sites: Amino acids 433-436;689-692

cAMP- and cGMP-dependent protein kinase phosphorylation:
Amino acids 232-235

Tyrosine kinase phosphorylation sites:

Amino acids 312-319;416-424

N-myristoylation site:

Amino acids 19-24;375-380;428-433;

429-434;432-437;517-522;574-579;

652-657;707-712

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66 Y Y N R S T S P W N L H R N E D P E R Y P S V I WE A K C R H L G C I N A D - G N V D Y H M N S V P

101 T H Q R S I S P W R Y R V D T D E D R Y P Q K L A F A E C L C R G C I D A R T G R E T A A L N S V R

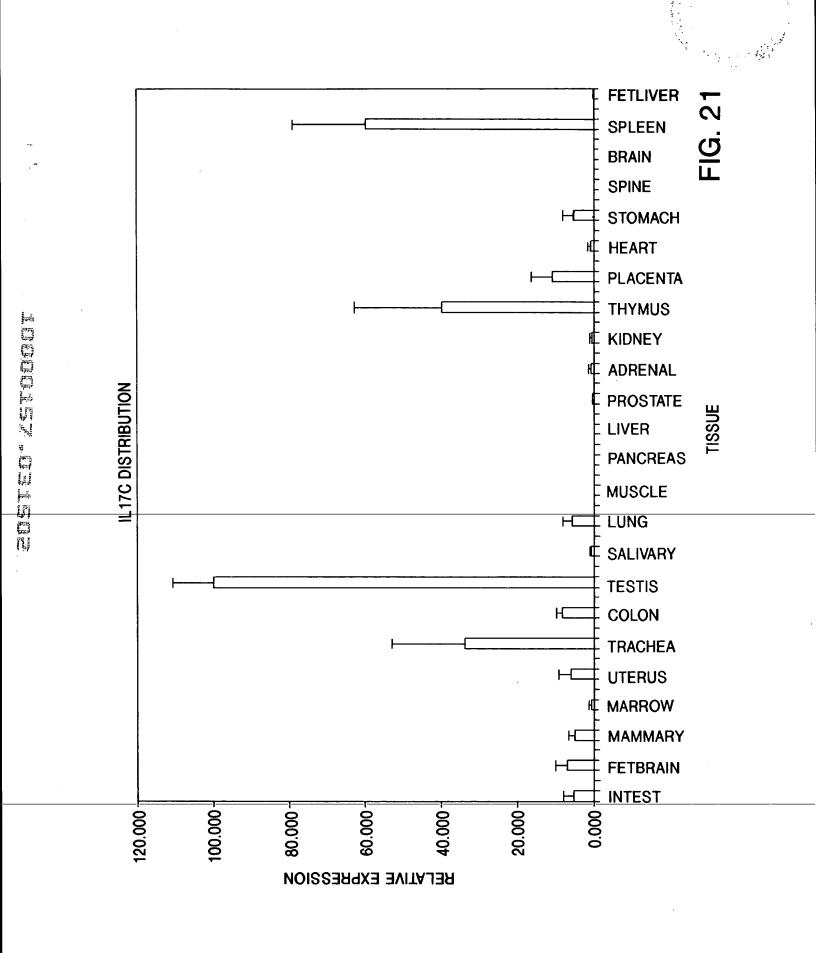
86 T N L R S V S P W A Y R I S Y D P A R Y P R Y L P E A Y C L C R G C L T G L F G E D V R F R S A P

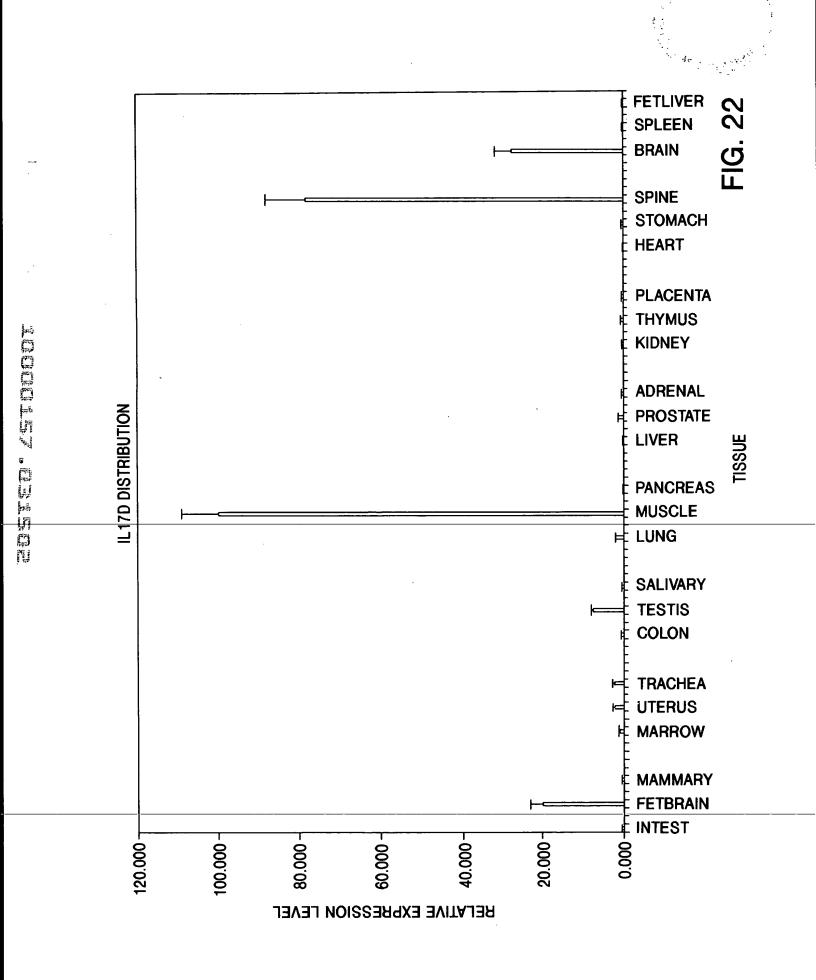
87 I N S R A I S P W R Y E L D R D L N R L P Q D L Y H A R C L C P H C V S L Q T G S H M D P R G N S E

74 I E S R S T S P W N Y T V T W D P N R Y P S E V V Q A Q C R N L G C I N A Q - G K E D I S M N S V P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         .....новсвузЕагЕкугутувстсутруіннуа
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ....GRSVYTEAYVTIPVGCTCVPEPEKDAD
                                                                                                    . . . . M L V A G F L L A L P P S W A A G A P R A G R R P A R P R G C A D R P
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            LLYHNOTVFYRRPCHGEK. GTHKGYCLERRLYRVSLACVCVRPRVB-
                                                                                                                                                                                                                                                             HQVPLDLVSRMKPYARMEEYERNIEEMVAQLRNSSELAQRKCEVNLQLW
LLARGAKWGQALPVALVSSLEAASHRGRHERPSATTQCPVLRPEEVLEA
                               ..... MD WP H N L L F L L T I S I F L G L G Q P R S P K S K R K G Q G R P G P L A P G
                                                              МТСГР G L L F L T W L H T C L A H H D P S L R G H P H S H G T P H C Y S A E E L | P | L | G | O | A | P P
                                                                                                                                     . . . . MRERPRLGEDSSLISLFLQVVAFLAMVMGTHTYS
                                                                                                                                                                                                                                                                                                                    EELLEOLY|G|RLAAGV|L|SAF|HHTLQLGPREQAR|N|ASCPAGGRPADRRFRP
                                                                                                                                                                      . . . . . . . . . . MTVKTLHGPAMVKYLLLSILGLAFLSEAAARKIPKV
. . MTPGKTSLVSLLLLLSLEAIVKAGI
                                                                                                                                                                                                                                                                                                                                                                WPSCCPSKGODTSEELLRWSTVPVPPLEPARPMAHPESCRAS---
                                                                                                                                                                                                                                                                                                                                                                                       HTFFQKPESCPPVPGGSMK DIGIINENORVS......
                                                                                                                                                                                                                                P G C P N S E D K N F P R T V M V N L N I H N R N T N T N P K
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          V Y M P T V V L R R T P A C A G
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Q Q E T L V V R R K
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                                                                                                                                                                                                                                                                                                   h-11.17c
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              h-IL17B
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          h-11.17D
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         h-11.17E
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            h-1117C
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       h-IL17F
                                            h-IL17B
                                                                                                       h-11.17b
                                                                                                                                        h-IL17E
                                                                                                                                                                         h-IL17F
                                                                                                                                                                                                                                                                       h-11.17B
                                                                                                                                                                                                                                                                                                                                    h-11.17b
                                                                                                                                                                                                                                                                                                                                                                   h-IL17E
                                                                                                                                                                                                                                                                                                                                                                                                  h-IL17F
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  h-11.17B
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                h-11.17D
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                h-IL17E
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               h-IL17F
                                                                         h-1117¢
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                h-IL17C
                                                                                                                                                                                                                                                                                                                                                                                                                                                                 h-IL17
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               h-IL17
                                                                                                                                                                                                                                       h-IL17
```

179

h-11.17D

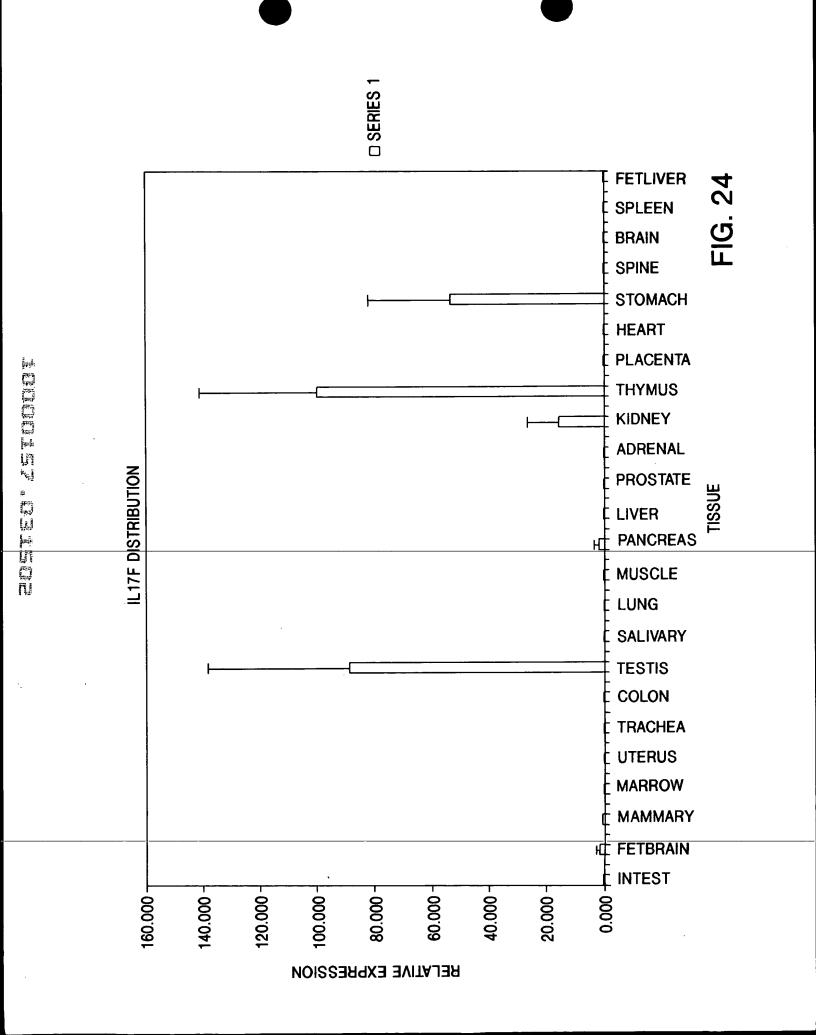


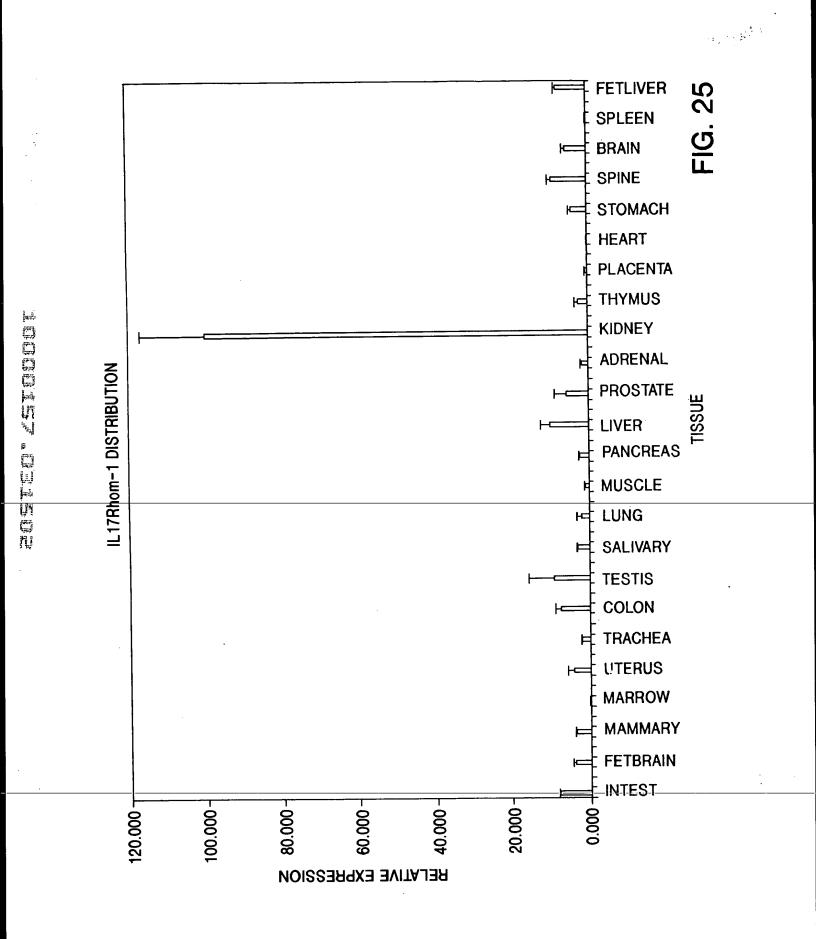


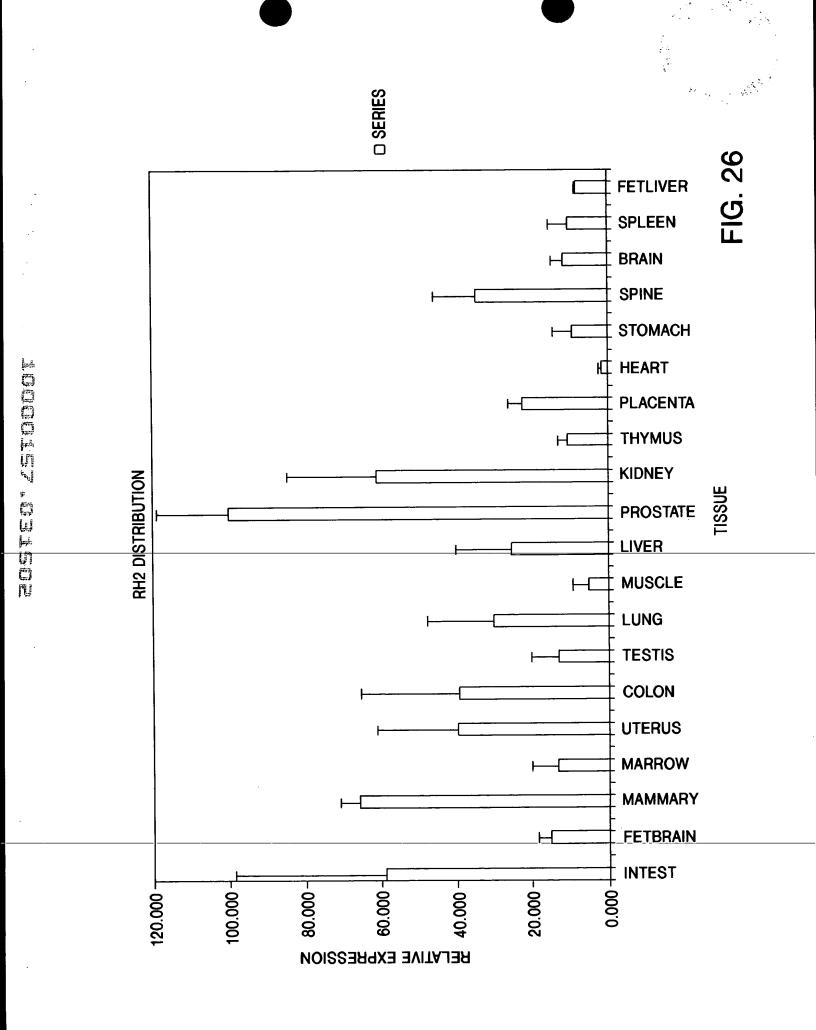
BRAIN
HEART
KIDNEY
LIVER
LUNG
COLON
MARROW
INTESTINE
SPLEEN
STOMACH
THYMUS
PROSTATE
MUSCLE
TESTIS
UTERUS
FETAL BRAIN
FETAL LIVER
SPINAL CHORD
PLACENTA
ADRENAL
ADRENAL
PANCREAS
SALIVARY
TRACHEA

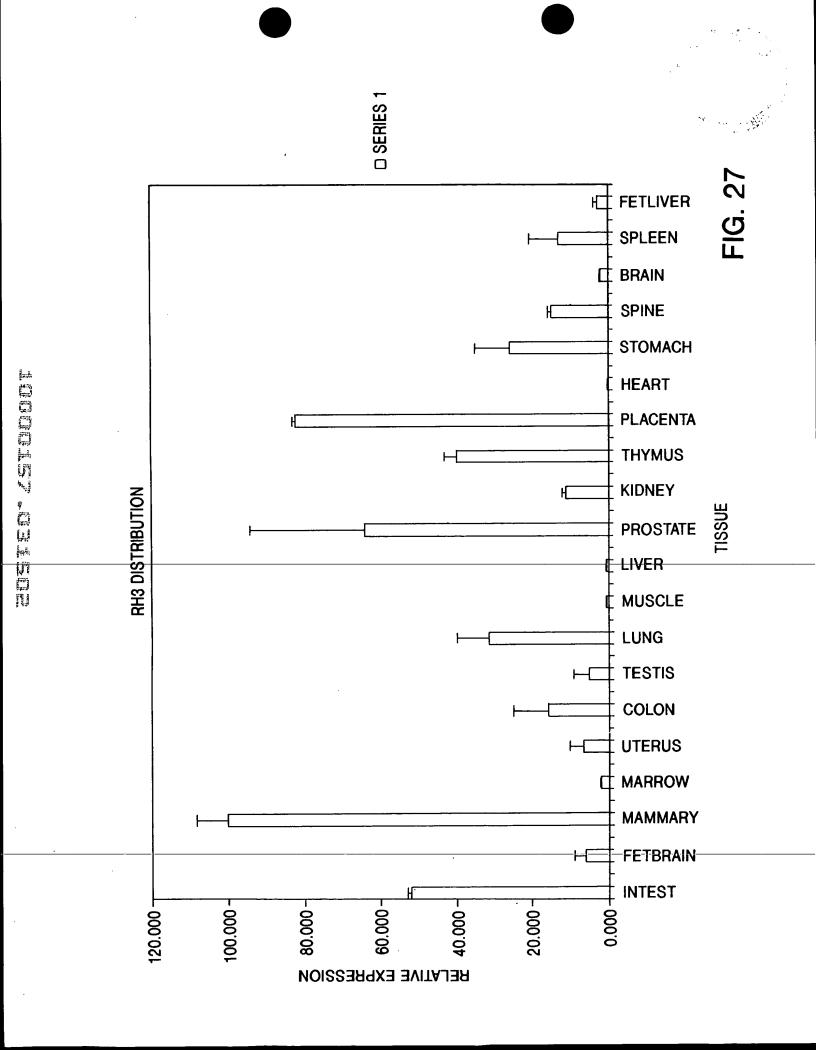
!

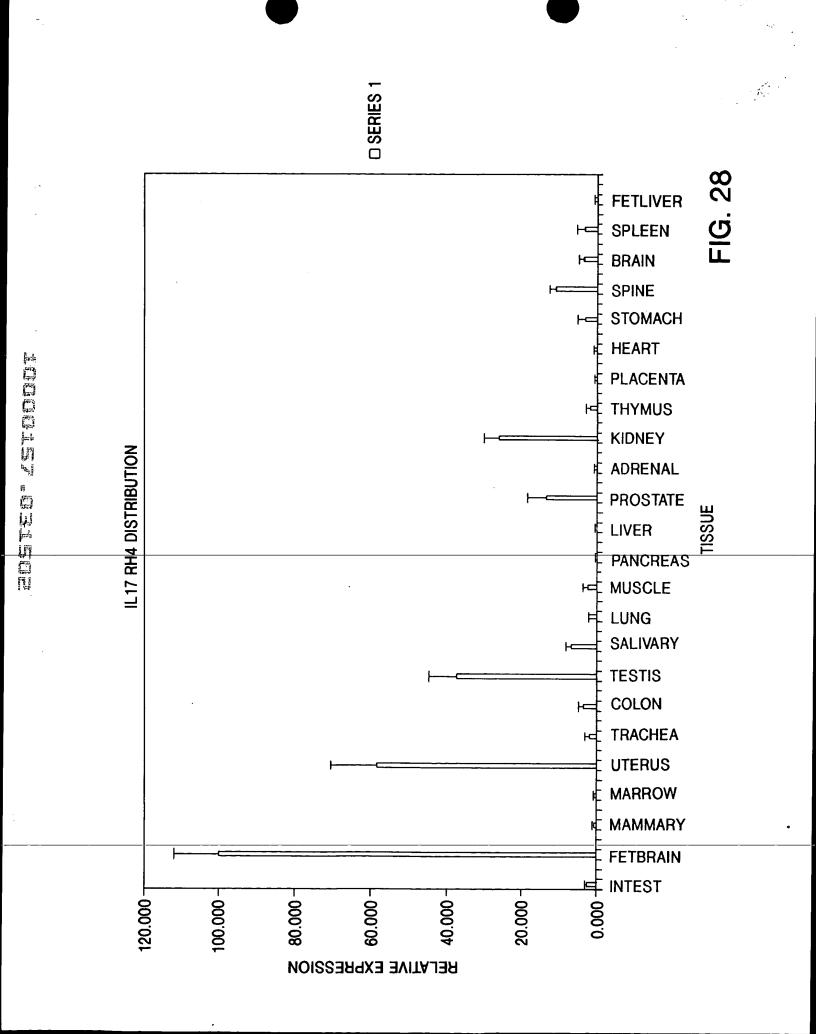
FIG. 23

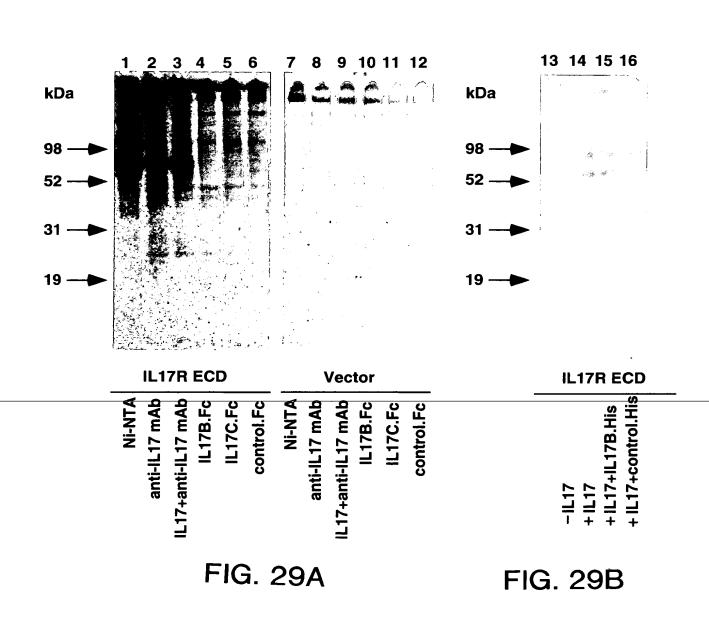












14.2

H Y E L D R D L N R E L L L L L L L L L L L L L L L L L L	THERETS AND THE TENT TO A PERSON AND THE TENT AND THE TEN	·
FIG. 30	134V H V A 196S V 175VM G	
<mark>АВВРОЗ В D G S G L Р ТР GA F A F H T E F П H V P V G C T C V L P Y R R P C H G E K G T H K G Y C L E R R L Y R V S L A C V C V R P</mark>		
RAIE PPHCPNSFALEKIILVSVGCTCVTP PRICPPP PRIGPCRORAVMETIAVGCTCIF -	10MNSVP 1QQE 1 LV L 39VSVPVFSQVPVRR	
RYRVDTDEDRYPOKLAFAECLCRGCIDARTG-RETA RYELDRDLNRLPODLYHARCLCPHCVSLOTGSHMDP	LEAUIHURSISPW - DGPLNSRAISPW	
NLHRNEDPERYPSVIWEAKCRHLGCINADG NVDY GYSINHDPSRIPVDLPEARCLCLGCVNPFTMQEDRS	- SSDYYNRSTSPW QLWMSNKRSLSPW	
G Q A L LPVALL V S S LELA A S H R G R H E R P S A T T Q C P V L R P E E SKGQ D T S E E L L R W S T V P V P P L E P A R P N R H P E S C R A S E	ള ഗ്രി	
N PGCPNSEDKNFPRTVMVNLNIHNRNTNTNPKR SRMKPYARMEEYERNIEEMVAQLRNSSELAQRKCEV		
FLIWEHTCLAHHDPSLRGHPHSHGTPHCYSAEELPLG LISLFEQVVAFLAMVMGTHTYSHWPSCCP		h-IL17C
<u>(ТSLVSLLL LLSLEA</u> IVK AGITIPR · · · · · · · · · · · · · · · · · · ·		h-IL17 :

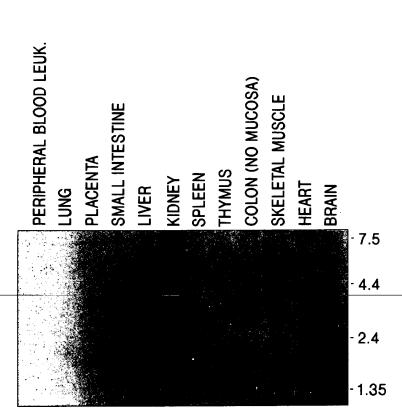


FIG. 31A

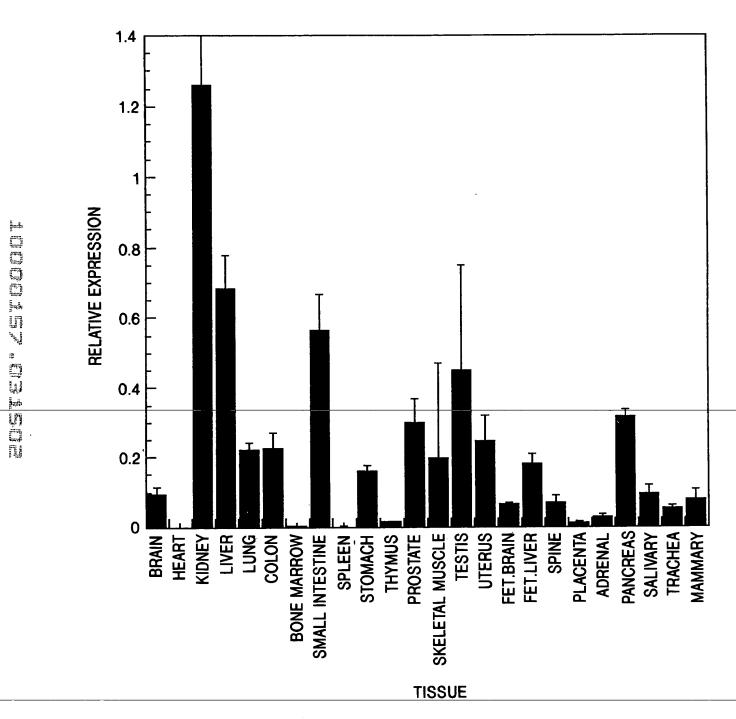


FIG. 31B

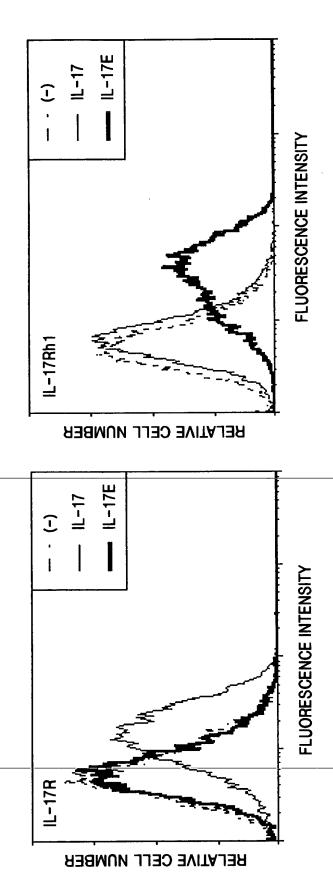


FIG. 32A

FIG. 32B

ķ.

FOLD INDUCTION LUCIFERASE ACTIVITY

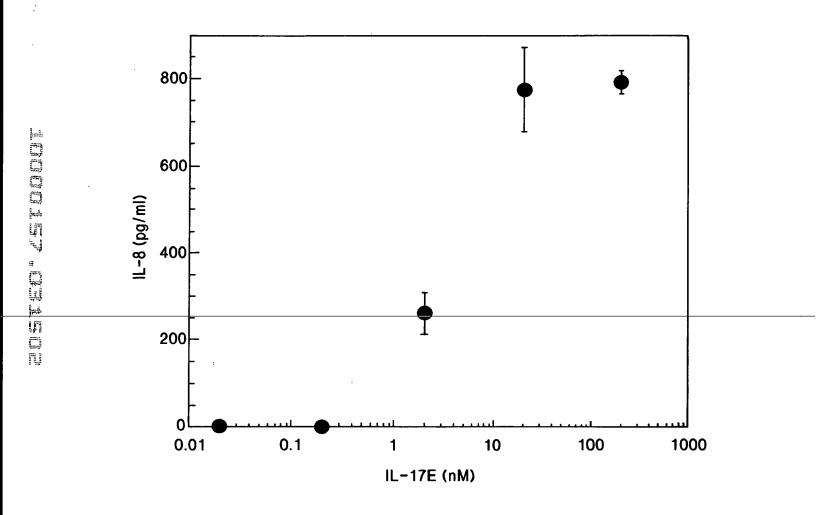


FIG. 34

IL-17 FAMILY OF CYTOKINES HAS COMPLEX PATTERN OF OVERLAPPING RECEPTOR-LIGAND SPECIFICITIES

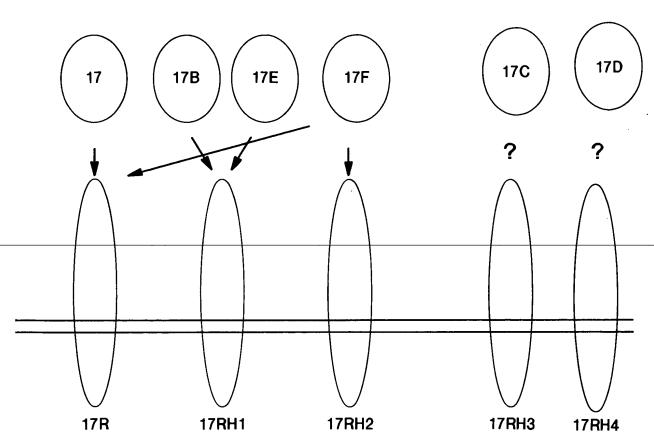
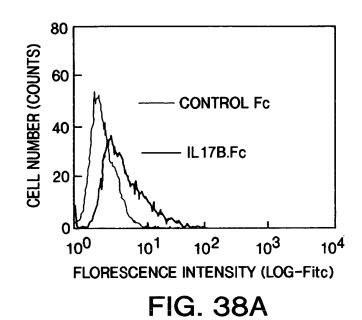


FIG. 35

HFF CELLS

160



SO CONTROL FC

40

100

101

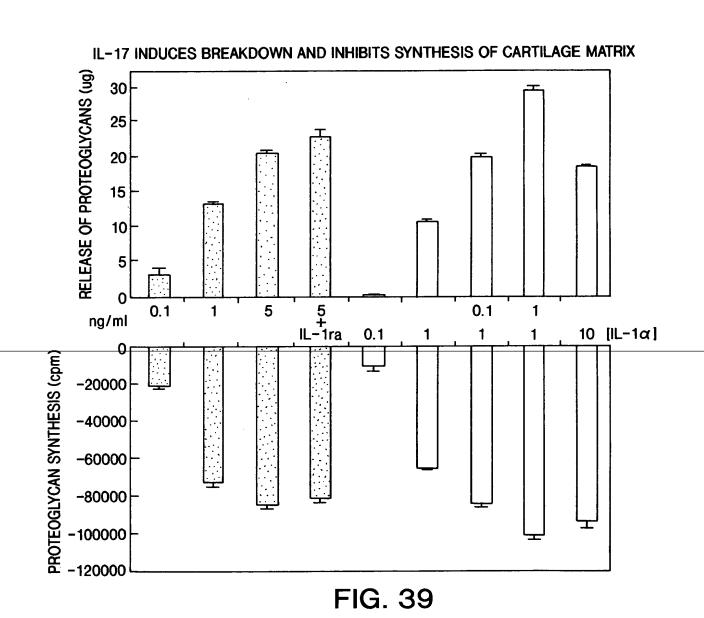
102

103

104

FLORESCENCE INTENSITY (LOG-Fitc)

FIG. 38B



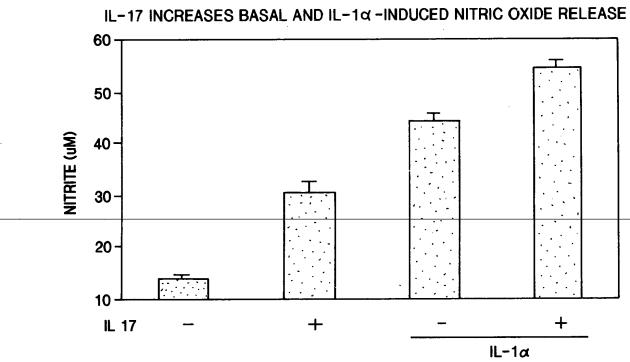
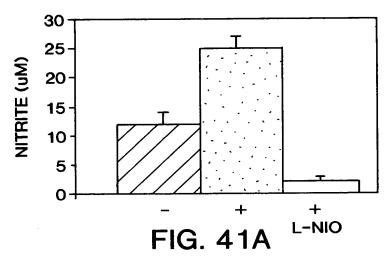
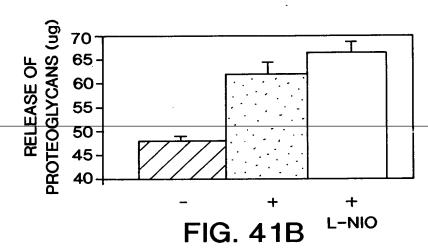
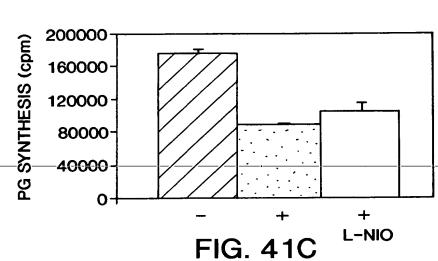


FIG. 40

INHIBITION OF NITRIC OXIDE RELEASE DOES NOT BLOCK THE DETRIMENTAL EFFECTS OF IL 17 ON MATRIX BREAKDOWN OR SYNTHESIS







INHIBITION OF NO RELEASE ENHANCES IL $1-\alpha$ -INDUCED MATRIX BREAKDOWN BUT NOT MATRIX SYNTHESIS

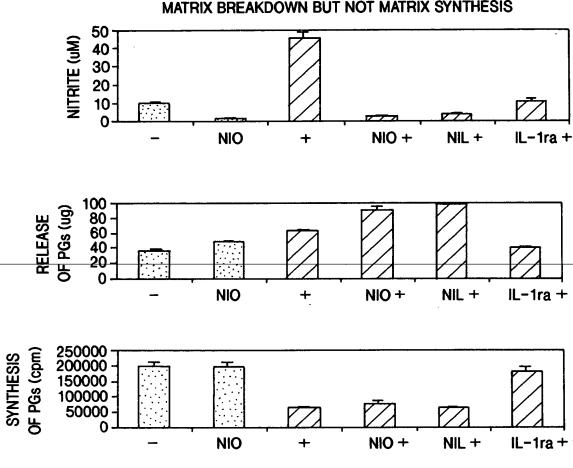


FIG. 42

IL-17C DETRIMENTAL EFFECTS ON ARTICULAR CARTILAGE

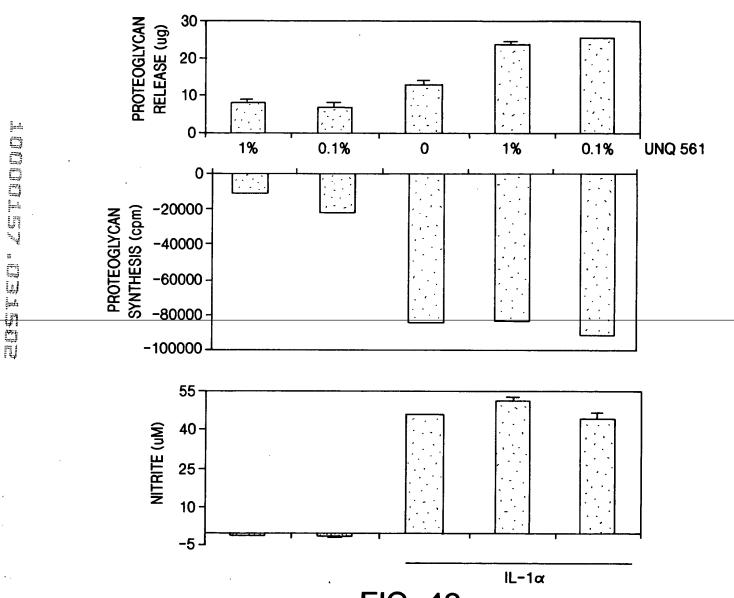
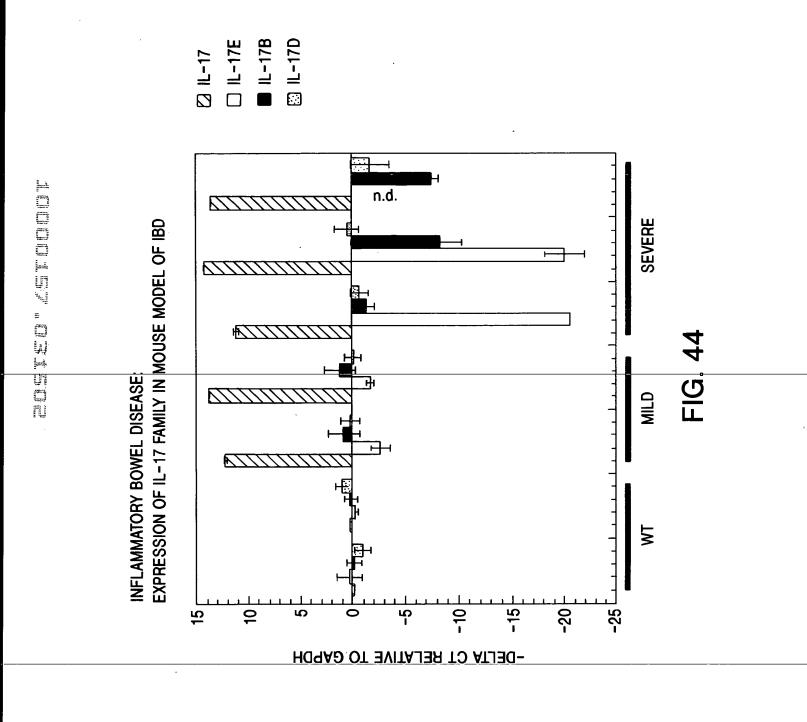
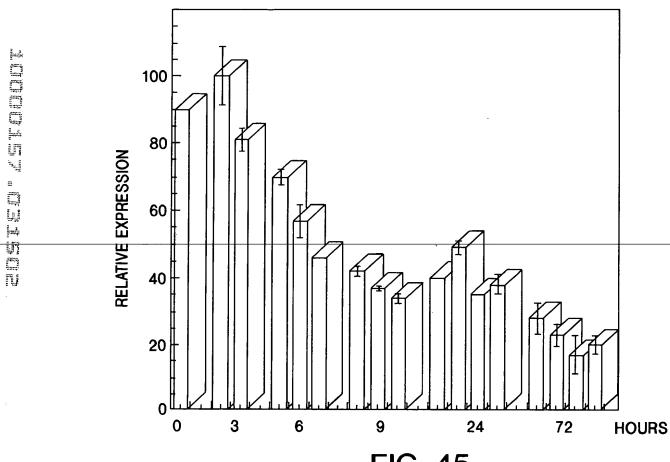
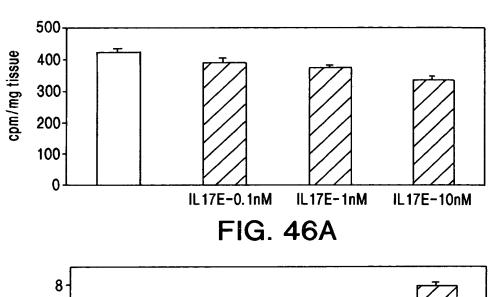


FIG. 43







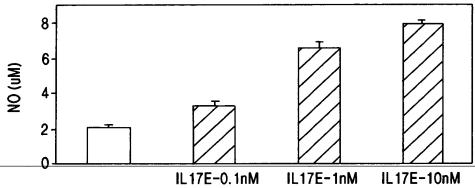


FIG. 46B

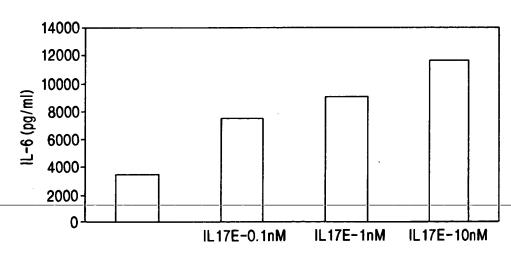
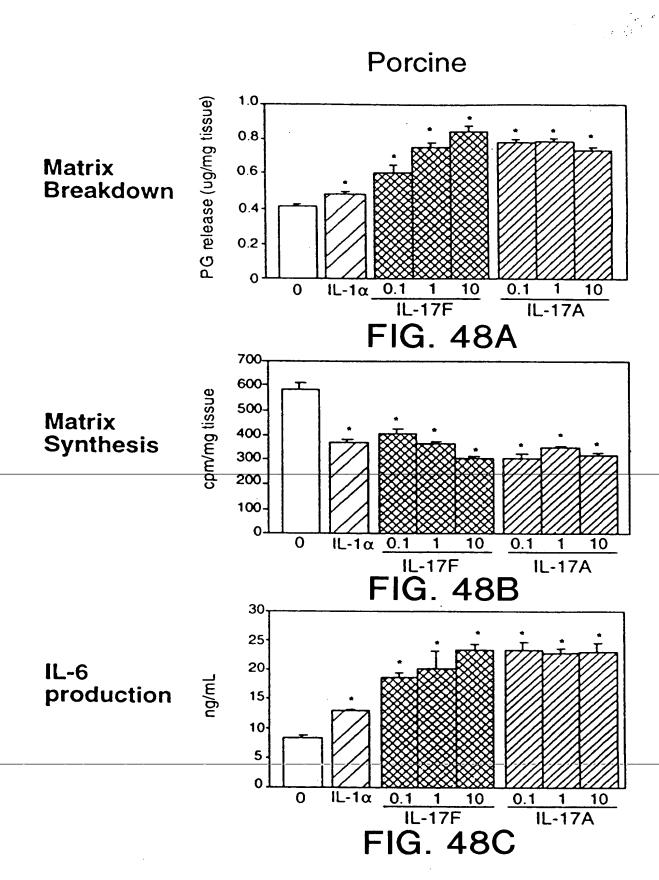
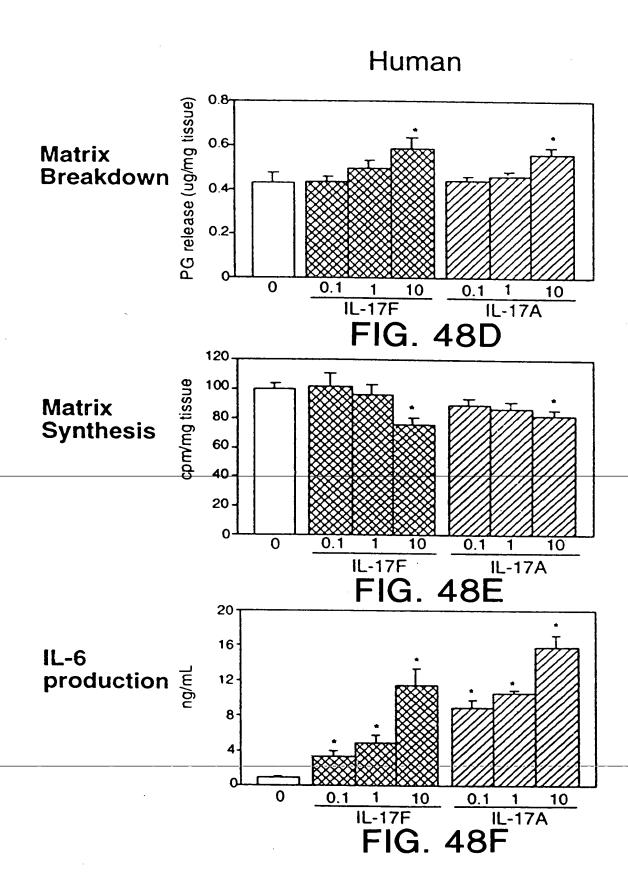


FIG. 46C





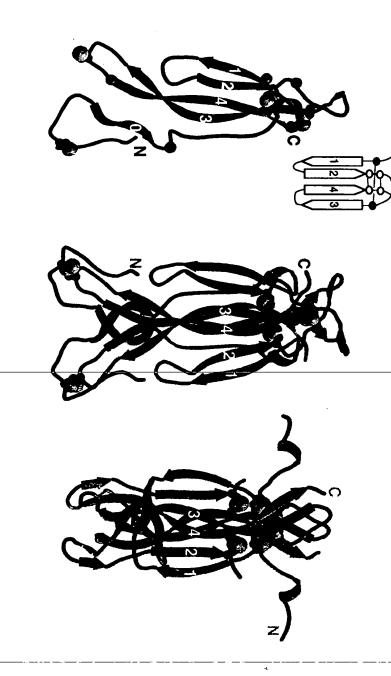


FIG. 49A

FIG. 49B

FIG. 49C

					*	•
IL-17F	•••••••	• • • • • • • • • • • • • • • • • • • •		RKIPKVG	HTFFQKPES	17
				IVRAG	ITIPRNP.G	14
TT -170	QPRS	PKSKRKGQGR	PGPLAPGPHQ	VPLDLVSRMK	PYARMEEYER	44
11-1/0	HHDPSLRGHP	HSHGTPH ≈YS	AEELPLGQAP	PHLLARGAKW	GQALPVALVS	50
TP-1/E	• • • • • • • • •	• • • • • • • • •	YS	HWPS PSKG	QDTSEELLRW	22
					·	
		4 0				
IL-17F	PPVPGG	SMKLDI	GIINENQRVS	MSRNIESRST	PWNYTVTWD	59
IL-17A	PNSEDKNFPR	TVMVNLNIHN	RNTNTN . PK	RESDYYMRST	DWNT.HDNPD	62
IL-17B	NIEEMVAQLR	NSSELAQR	KEVNL	OLWMSNKRSL	PWGYSTNHD	ΩΩ
IL-17C	SLEAASHRGR	HERPSATT	O PVLRPEEV	LEADTHORST	DWBVBVDVID	90
IL-17E	STVPVPPLEP	ARPNRHPE	Serase	.DGPLNSRAI	PWRYELDRD	65
		2				
TT17P	DNDVDCBIRG		# 1 : 00			
TT173	PNRYPSEVVQ	WO SKUTG STU	AQGKEDIS	MN VPI.QQE	TLVVRRKHQG	106
II-178. II-178	PERYPSVIWE	WY KUITGEIN	A. DGNVDYH	MN VPI.QQE	ILVLRREPPH	109
TT176	PSRIPVOLPE	WK TI TO VN	PF.TMQEDRS	MV VPV.FSQ	VPVRRRL	133
111-17C	EDRYPOKLAF	WE TO KO TD	AR. TGRETAA	LN VRL.LQS	LLVLRRRP	144
TU-T/E	LNRLPQDLYH	WKSTOPHIOS	LQTGSHMDPR	GN ELLYHNQ	TVFYRRP	112
						
			A SON STA	6-		
	*	Salah M.	स्यो स्ट १८ ३ (३८५) महिन्द्यो (१५)	W		
IL-17F	esv	SFOLER	VLVTVG	L-VIIPVIHHVO		133
IL-17A	EPN	SFRLEK	IL.VSVG	COMPIVATOR	• • •	136
IL-17B	PPPPRTGP.	RORA	VMETIAVGET	CIF	• • •	160
		PGAFAFHTEF	IHVPVG	W. LPRSVAA	AT.R	184
	HGEKGTHKG	YLER	RLYRVSLACV	AVRPRVMG	- secol	145
						49:1

FIG. 50

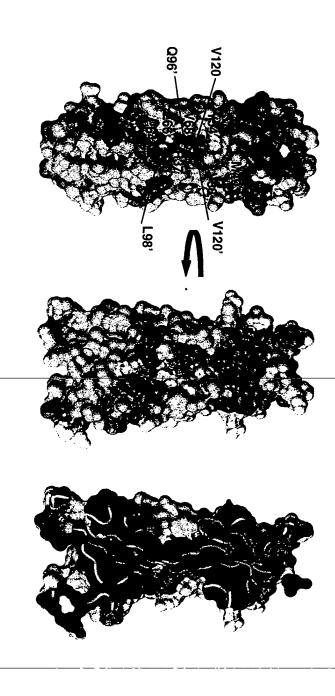


FIG. 51A

FIG. 51B

FIG. 51C

FIG. 52C

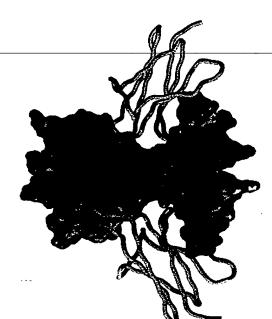


FIG. 52A

FIG. 52B

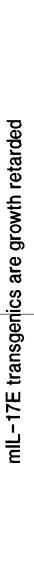
IL-17E is highly conserved between human and mouse

SHWPSCCPS ---VAFLAMIVGTHTWSLRIQEGCISHL SLISLFLQVVAFLAMVMGTHT ß GED MRERPRL mlL-17E NL-17E 35 KEQEPPEEWLKWSSASVSPPEPLSHTHHAESCRASKDGPLNSRAISPWSY 43 KGQDTSEELLRWSTVPVPPLEPARPNRHPESCRASEDGPLNSRAISPWRY mL-17E NL-17E

85 ELDRDLORYPYDLYHARCLCPHCVSLQTGSHMDPLGNSIVPLYHNQTV FYR 93 ELDRDLNRLPQDLYHARCLCPHCVSLQTGSHMDPRGNSELLYHNQTVFYR ml-17E NL-17E

KPCHGEE GTHR RIYCLERKLIYKV SLACVCV KPKVMA G THK GY CLERRLY RV SLACVCVRPRVMG RРСИ G EK 135 143 mL-17E **ML-17E**

FIG. 53



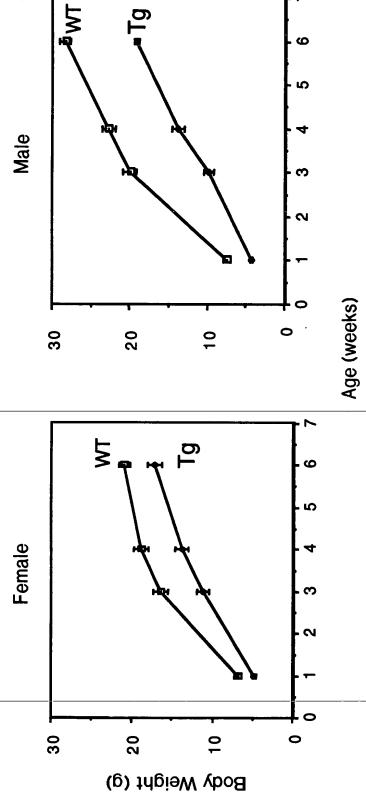


FIG. 55

IL-17E transgenics are jaundiced by 6 weeks of age

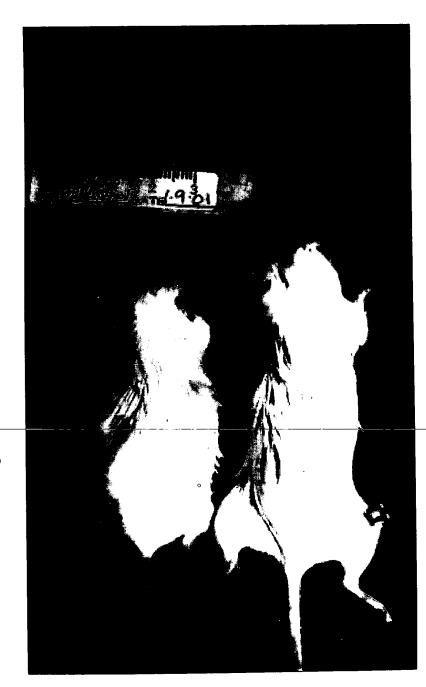
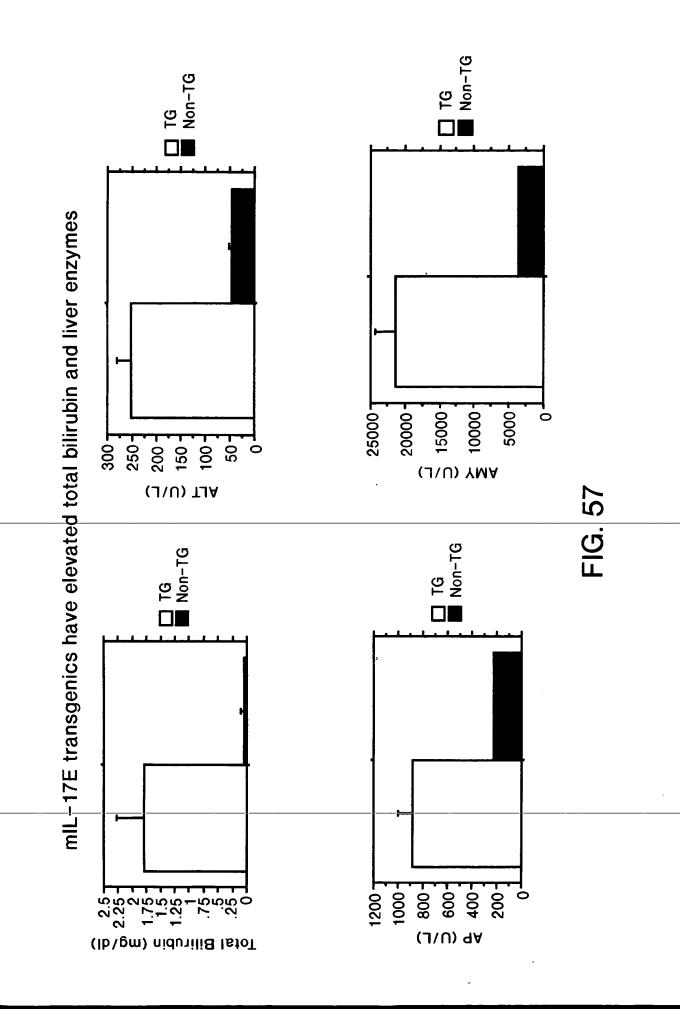
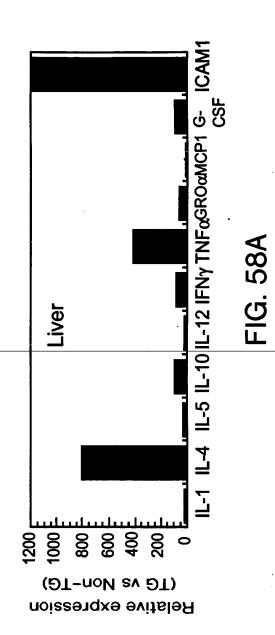


FIG. 56





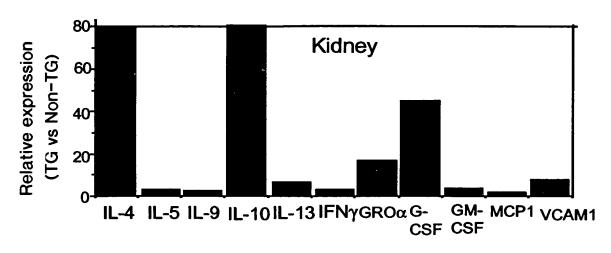


FIG. 58B

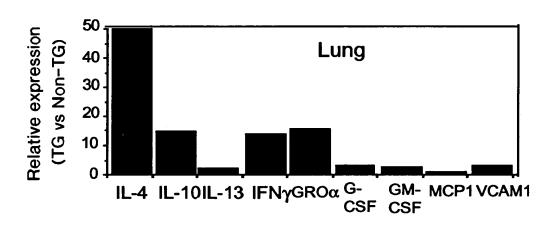


FIG. 58C

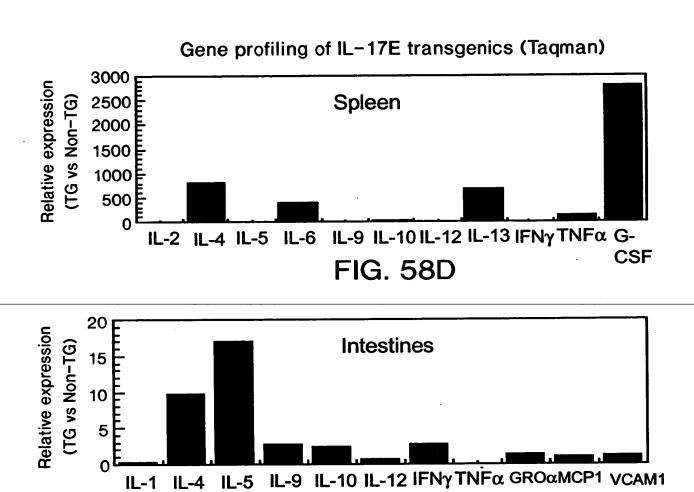


FIG. 58E



Elevated serum IL-5, IL-13 and TNF α in mIL-17E transgenics

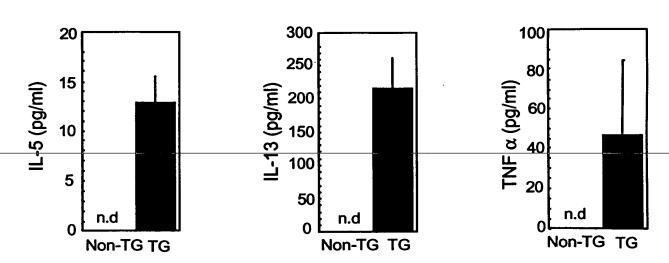


FIG. 60

Serum IgE and IgG1, but not IgG2a is elevated in mIL-17E transgenics

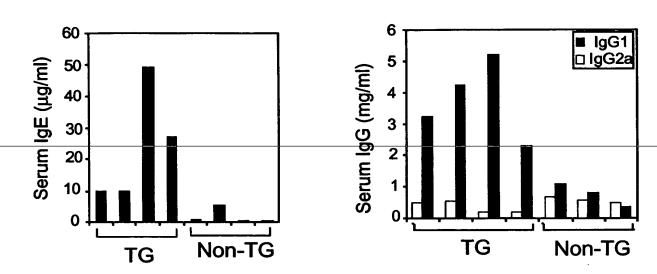
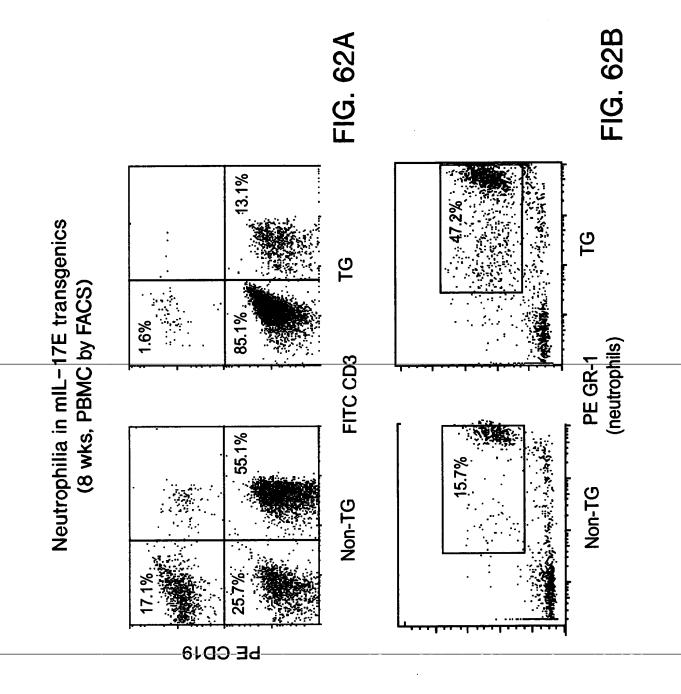


FIG. 61



Neutrophilia and eosinophilia in mIL-17E transgenics (hematology)

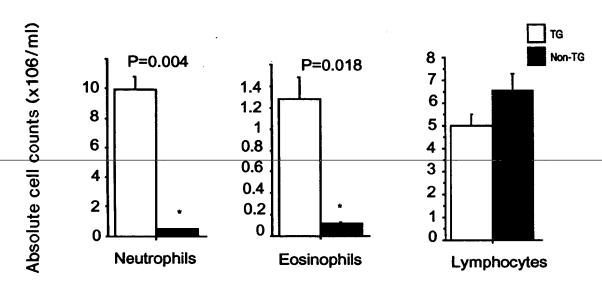


FIG. 63

G-CSF is elevated in mIL-17E transgenics

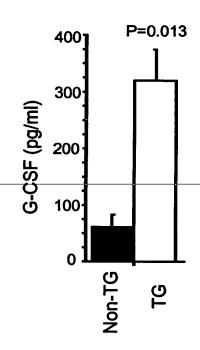


FIG. 64

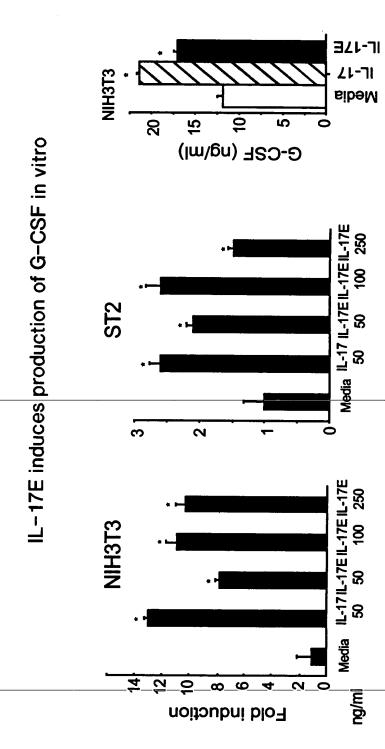


FIG. 65

*P<0.05

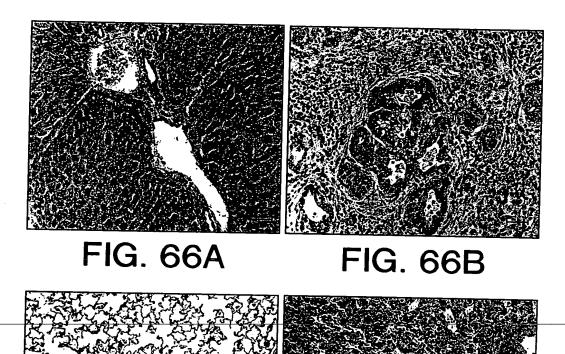


FIG. 66C

FIG.66D